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GRANT NUMBER DAMD17-94-J-4421

TITLE: Breast Cancer Research Training Grant

PRINCIPAL INVESTIGATOR: Adrianne E. Rogers, M.D.

CONTRACTING ORGANIZATION: Boston University Medical Campus

Boston, Massachusetts 02118

REPORT DATE: October 1996

TYPE OF REPORT: Annual

PREPARED FOR: Commander

U.S. Army Medical Research and Materiel Command Fort Detrick, Frederick, Maryland 21702-5012

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FOREWORD

Opinions, interpretations, conclusions and recommendations are those of the author and are not necessarily endorsed by the U.S. Army.

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MH Where material from documents designated for limited distribution is quoted, permission has been obtained to use the material.

Citations of commercial organizations and trade names in this report do not constitute an official Department of Army endorsement or approval of the products or services of these organizations.

In conducting research using animals, the investigator(s) adhered to the "Guide for the Care and Use of Laboratory Animals," prepared by the Committee on Care and use of Laboratory Animals of the Institute of Laboratory Resources, national Research Council (NIH Publication No. 86-23, Revised 1985).

NH For the protection of human subjects, the investigator(s) adhered to policies of applicable Federal Law 45 CFR 46.

In conducting research utilizing recombinant DNA technology, the investigator(s) adhered to current guidelines promulgated by the National Institutes of Health.

In the conduct of research utilizing recombinant DNA, the investigator(s) adhered to the NIH Guidelines for Research Involving Recombinant DNA Molecules.

In the conduct of research involving hazardous organisms, the investigator(s) adhered to the CDC-NIH Guide for Biosafety in Microbiological and Biomedical Laboratories.

PI - Signature

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BREAST CANCER RESEARCH TRAINING PROGRAM (BCRTP)

5. INTRODUCTION

Information about breast cancer is increasing rapidly from many sources, but much of it is inconsistent and controversial because of the complexity of the disease and its causes. Difficulties arise also in integration of knowledge and understanding and in extrapolating results between and within scientific disciplines. The Program in Research on Women's Health at Boston University Medical Center (BUMC) has been developed to address some of these problems by serving as an integrating and promoting force for research in women's diseases with a major effort focused in breast cancer research. Through the research and teaching collaborative efforts already in place and with the stimulus of this Training Program the work of investigators at BUMC is being brought to bear on training students to conduct research on breast cancer.

The **PURPOSE** of the Program is to: 1) establish a formal multidisciplinary research and academic training program in breast cancer biology and epidemiology; 2) produce graduates in one discipline (Pathology, Epidemiology, Microbiology) who have an understanding of the other disciplines and who can perform collaborative, multidisciplinary research in the etiology, prevention and therapy of the disease; 3) provide training in cell and molecular biology, experimental pathology, carcinogenesis, epidemiology and biostatistics, immunology, toxicology, and nutrition that will permit trainees to explore: a) basic breast cancer cell processes and interactions, including oncogene regulation, cell signalling, and genetic considerations in design of therapeutic agents; b) questions about etiology, prevention and therapy of breast cancer in laboratory animals and human populations; and c) the integration of knowledge derived from the different approaches; 4) maintain and increase collaborative research in breast cancer and closely related areas among faculty and trainees; 5) provide attractive opportunities for all students and, specifically, for women and underrepresented minorities to pursue careers in breast cancer research. We are promoting the development of young investigators who will have a broad, multidisciplinary background in breast cancer biology and epidemiology and intensive training in a specialized research area and who can perform significant research using advanced concepts and techniques and communicate research accomplishments effectively. They will be a resource to meet future personnel requirements for breast cancer research. The **METHODS** are as follows. Doctoral students committed to cancer research with an interest in breast cancer research are admitted to the Departments of Pathology

and Laboratory Medicine, Epidemiology or Microbiology and follow a curriculum specifically designed for this Training Program. BCRTP ensures that each predoctoral student: 1) participates in an appropriate, integrated curriculum focused on breast cancer; 2) has an advisory committee, composed of basic science and epidemiology faculty members with expertise in or closely related to breast cancer research; 3) participates actively in seminars, and local, regional and national meetings in addition to informal research meetings at the school.

The students are closely integrated into the <u>Breast Cancer Working Group</u> in the Program in Research on Women's Health. The Group comprises over 50 members in multidisciplinary teams collaborating in breast cancer research and developing new research strategies. The Group stimulates research interactions by providing teaching and discussion of clinical and research topics at monthly meetings. The BCRTP is significantly extending and supplementing the doctoral programs in the participating departments. The BCRTP is directed by Adrianne E. Rogers, MD, and Theodore Colton, DSc, with extensive input from Gail Sonenshein, PhD, and Marianne Prout, MD, who direct the Research on Women's Health Program.

Drs. Rogers and Colton meet at least twice each term and take responsibility for all decisions on student admission, performance and training with substantial input from the faculty Trainers and from the Trainees themselves. Because of the relatively small size of the Program and the extensive interactions with faculty in the Women's Health Program, the Admissions and Performance, Recruitment and Seminar Committees proposed have not been needed and, therefore, have not been formally set up.

The trainees are supported annually by the BCRTP and BUSM and BUSPH. The two schools have supplemented the BCRTG so that each student receives a stipend of \$14,000-\$15,000 per year and tuition for 20-24 credits per year as needed. They have been admitted to the Program via the Pathology and Laboratory Medicine or the Biostatistics and Epidemiology Departments. In the beginning of each academic year all new and returning trainees have met with Drs. Rogers and Colton to review the Program and its requirements and opportunities, and to answer questions and plan each student's curriculum. Trainees are encouraged to consult any of the participating faculty for general advice or further discussion of their research interests, and are directed to appropriate faculty by Drs. Rogers and Colton. Trainees have additional contact with faculty members in courses and seminars.

Drs. Rogers and Colton have met also with each trainee at the end of each semester to discuss trainees' academic performance, to obtain feedback about the program and to advise the trainee on choice of courses and lab rotations. When the

students move into their dissertation research, their Faculty Trainer becomes their major adviser, but Drs. Rogers and Colton continue to meet individually with them at least once a year. After passing qualifying exams, trainees' research progress will be reviewed on a regular basis by their dissertation committees, composed of three members from the home department Training Faculty and one member each from the other two departments' Training Faculty. Under appropriate circumstances (need for a particular expertise), one of the committee positions may be filled by non-Trainer faculty from inside or outside BUMC. These committees will meet with the trainee at least twice a year, usually starting within six months of the qualifying exam when the student presents her or his thesis research proposal. Finally, the thesis committee will serve as the examining committee for the thesis defense.

6. **BODY**

The six students are:

1994: Yvette Cozier (BA, Liberal Arts, Harvard Extension School, 1987; MPH, BUSPH, 1994) was admitted to the Biostatistics and Epidemiology DSc. program. She had extensive laboratory experience in Hematology and in Microbiology (1982-1994), strong letters from faculty and a 3.5 GPA at BUSPH. She was particulary interested in Dr. Lynn Rosenberg's epidemiological studies in breast cancer and other diseases in black women, and has joined Dr. Rosenberg's group for her dissertation research. She is currently completing her SPH course requirements and plans to take the qualifying exmaination in summer, 1997. Yvette is an African-American.

Laurie Hafer (BS, Microbiology, Penn. State Univ., 1989) was admitted to the Pathology and Laboratory Medicine Program. She had extensive clinical and research experience in the Immunohistochemistry laboratory at the College of Medicine-University Hospital, Hershey, PA, where she was in charge of research and development with a major focus on breast cancer studies. She had very strong letters from faculty who had supervised and worked with her. She entered intending to pursue interests in immunology and breast cancer but has become interested in hormone receptor cellular and molecular biology and is developing her research proposal in that area with Drs. Rogers and Traish. She worked extensively with Drs. Rogers and Delas Morenas last year in histological and immunohistochemical studies of rat mammary tumors and on image analysis of ER & PCNA in them and with Dr. Carol Rosenberg in molecular studies of breast cancer. The specific subject was the methylation status of the TGFb receptor II promoter in breast cancer and other cell lines. She has completed her course requirements and passed the qualifying examination in June, 1996. She participated in the AACR Keystone

workshop on Histopathobiology of Neoplasia in July, 1996, and presented a poster there.

1995. Sylvia Marecki (BS, Microbiology, Univ. N.H., 1995) was admitted by the Pathology and Laboratory Medicine PhD Program and is in the immunology track. She had significant undergraduate research experience and was awarded two competitive research grants in addition to a four-year scholarship. She had very strong letters from her research adviser and other faculty. As we discussed research opportunities in Immunology, her interests shifted to cancer research from her initial focus on bacteriology, which was the subject of her undergraduate research. She is completing her course requirements and will take the qualifying exam in June, 1997. Sylvia did an excellent job in her first lab rotation with Dr. Rogers in the rat DMBA carcinogenesis study, has completed two additional rotations (Dr. Beller: II-1 defect in autoimmunity; Dr. Wexler: B-cell, costimulation) and is completing a rotation with Dr. Fenton in molecular biological studies of cellular immune responses; she is well-oriented to breast cancer research and plans to work with Dr. Fenton in studies of macrophage function. She was a co-author on Laurie Hafer's poster at the AACR workshop.

Paul Johansen (formerly Mange) (BS Biology, Yale, 1988) was admitted to the Biostatistics and Epidemiology PhD program. This program differs from the DSc program in the SPH in being a more extensive joint program with the Mathematics Dept. on the Charles River Campus of BU and in requiring a more sophisticated mathematics and biostatistics curriculum and dissertation. After completing 1 1/2 years of medical school at Univ. Mass, Paul left to pursue interests in math and statistics and worked in biomedical applications of these areas as Sr. Research Analyst in Psychiatric Epidemiology at the Mass. General Hospital. He had excellent letters from faculty and colleagues. Paul is completing courses required for his program and is teaching in an elementary biostatistics course. He completed the practical carcinogenesis course and is contributing valuable statistical questions and insights to current discussions of Laurie Hafer's studies described above.

Thus a major goal of the program (training and encouraging students in interdisciplinary, collaborative research) is being achieved.

1996 Jackie Ashba (BA, Biology & Economics, Clark University, 1989; MPH, BUSPH, 1992; MA, BUSM, Medical Sciences, 1994) was admitted to the Epidemiology DSc program. Her Masters degree research included studies of both biological and epidemiological aspects of breast cancer. Her academic record here and her letters of recommendation are excellent. She has begun courses and is working in a DMBA mammary carcinogenesis project with Dr. Rogers to begin her practical experience in this field.

Ingrid Gherson (BS, Biology, Binghamton University SUNY) was admitted to the Dept. of Pathology and Laboratory Medicine. Her excellent undergraduate record (GPA 3.5) and GRE scores, strong letters of recommendation and significant technical experience in a pathology laboratory all are evidence of her potential for success in the program. She is working in the DMBA carcinogenesis project with Dr. Rogers and taking her initial courses in biochemistry and epidemiology.

The BCRTP students participate actively in the practical course in the setting up and running of DMBA mammary tumorigenesis studies under Dr. Rogers' direction. They learn basic methods for such studies, have participated in feeding, weighing and observing the animals and in performing necropsies for examination of tumors and collection of tissues for histological, endocrine and molecular studies. Members of the Pathology faculty introduce them to clinical studies of breast cancer. The subjects specifically covered are: Dr. De las Morenas: basics of breast cancer pathology; Dr. Burke: basics of image analysis, focused on estrogen receptor assay; Dr. Yang and one of the students, Laurie Hafer: basics of immunohistochemistry staining and interpretation; Dr. Rogers: basics of the histopathology of rat mammary gland tumors and discussion of recent research papers.

The students participated in both the Research in Women's Health and the Pathology seminar series and in Breast Cancer Working Group meetings in addition to a variety of other seminars in the two schools and in the Mass. Breast Cancer Research Program. This year the BCRTG hosted two special seminars (Drs. Robert Clarke and Michael Conner). The students had a luncheon and discussion with each of them before the seminar, and also joined other students for discussion with Dr. Lawrence Grossman (announcements attached).

7. CONCLUSIONS

The Program is actively recruiting, attracting and retaining excellent students from diverse backgrounds to focus on breast cancer research. The students are a cohesive group who study and work together well. They interact extensively with Drs. Rogers & Colton and with other students and faculty working in breast cancer research and working in clinical settings with breast cancer. They are doing well in course work and in their initial research. The interdisciplinary focus in strong, fostered by the practical course, the required epidemiology and pathology courses, seminars, and frequent formal and informal meetings of the students with Drs. Rogers and Colton. The students are progressing as expected (or more rapidly than expected) through their course work and into research, a commendable result.

8. References

None

APPENDICES

September, 1995: L-R: Paul (Mange) Johansen, Sylvia Marecki, Dr. Colton, Yvette Cozier, Dr. Rogers, Laurie Hafer



September, 1996: L-R: Laurie Hafer, Dr. Rogers, Sylvia Marecki, Paul(Mange) Johansen, Yvette Cozier, Jackie Ashba, Dr. Colton, Ingrid Gherson

Appendix b

Commonwealth of Massachusetts The Trial Court

Division	Probate and Family Court Department	Docket No. 76 C 2739 C
PAVI	Change of Name(s) FRANKLIN MANGE	
(Street and No.)	UE SUMERVILLE MIDD (City or Town)	<u> したら巨大 リフリイヤーリリク</u> (County) (Zip)
Occupation Graduate	student in Biostatistics Bos nuary 24 1966 in Northampt	ston University
Date and Place of Birth La	nuary 24 1966 in Northampt	on, massachusetts
Name of Spouse		
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Have parties ever changed their	names prior to this petition? <u>N U</u> If answer	is yes, explain
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Child		
Date March 1 1996	Paul F	ranklin Manaz
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	mother as next fri	• •
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his/her/their legal name(8).	1 1006	E Mr. Johnson
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CJ-P 27 (8/88)

Middlesex S.S.

Probate Court
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Register

Histopathobiology of Neoplasia

"Black Tea and Mammary Carcinogenesis in Rats", <u>Laurie J. Hafer</u>, Yvette S. Iskander, Sylvia Marecki, Adrianne E. Rogers, MD, Boston University School of Medicine and Mallory Institute of Pathology, Department of Pathology and Laboratory Medicine

Evidence of anticarcinogenic effects of diet components has been reported. Extracts of green tea in particular have been shown to prevent or reduce carcinogenicity in the skin, lung, esophagus, forestomach and duodenum of laboratory rodents. Studies of black tea extracts have yielded similar results, but have been less extensive. Breast cancer, a leading cause of morbidity and mortality among women, has also been shown to be susceptible to dietary intake in animal models, particularly intake of unsaturated fats, Vitamin A and selenium. Thus, the question of the effect of tea on mammary carcinogenesis was posed. Since the majority of the tea drinkers in the west consume black tea, the decision was made to examine its effect in rats. Female Sprague-Dawley rats were randomized into either deionized water or tea drinking groups and fed an AIN-76A diet. At 55 days of age, the rats were given a single dose of 7,12dimethylbenz(a)anthracene (DMBA) by gastric gavage. Rats were palpated twice weekly for tumors and killed by CO₂ inhalation when the tumors reached 3 cm in diameter. The remaining animals were killed 16 weeks post-DMBA administration. Tumor latencies, incidences and numbers were determined for each of the groups. Two experiments were performed using identical protocols except the dose of DMBA which was 25 mg/kg (Expt #1) or 15 mg/kg (Expt #2). Rats were given 1.25% or 2.5% tea extract.

There was no statistically significant effect of black tea on mammary tumor development in either experiment. Ongoing studies are examining the morphological, biochemical, and molecular changes in tissue samples obtained from these animals.

Results:

Results:			
	<u>Tumor</u>	<u>Tumor</u>	<u>Tumor</u>
	Latency (d)	Incidence (%)	Number per TBA ^a
Expt#1:			
Control	80	60	2.9 + 1.7
1.25% Tea	71	75	3.4+2.3
Control	76	62	2.8+1.9
2.5% Tea	72	68	3.2+2.1
Expt#2:	·		
Control	101	62	2.0+1.2
1.25% Tea	90	40	1.8 + 1.2
Control	103	45	1.7 + 0.8
2.5% Tea	98	40	2.0+1.6

^aTBA=Tumor bearing animal

This research is supported by the Tea Trade Health Research Association and USAMRDC Breast Cancer Research Training Grant.

BOSTON UNIVERSITY SCHOOL OF MEDICINE/SCHOOL OF PUBLIC HEALTH . THE UNIVERSITY HOSPITAL . BOSTON UNIVERSITY GOLDMAN SCHOOL OF GRADUATE DENTISTRY



Department of Pathology and Laboratory Medicine

80 East Concord Street Boston, Massachusetts 02118-2394 TEL: 617 638-4500 FAX: 617 638-4085

Memorandum

DATE: February 14, 1996

TO: Breast Cancer Research Trainees and Mika Sovak, Dong Kim, Anthony

Trombino, Koren Mann, Yvette Iskander and Alex Shneider

FROM: Adrianne E. Rogers, M.D.

RE: Spring Seminar Speakers

The Training Grant is sponsoring two speakers this term in conjunction with the Program in Research on Women's Health. You are invited to have lunch with them and to meet with them for informal discussion. If there are other students in your labs who would be interested to attend, please invite them and let me know.

The speakers are:

Thursday, March 21:

Robert Clarke, Ph.D.

Associate Professor, Lombardi Cancer Center and Dept. of Physiology & Biophysics, Georgetown University

Seminar title: Estrogen regulated genes and acquired endocrine resistance in breast cancer.

Thursday, May 23:

Michael Conner, DVM

Seminar title: Inhibition of farnesyltransferase induces regression of mammary and salivary carcinomas in *ras* transgenic mice.

On March 21, we'll have lunch with Dr. Clarke, 12:00-1:00 in Hiebert Lounge, 14th floor, and then he'll stay on for a discussion with you until about 2:00. His seminar is at 4:00.

On May 23, Dr. Conner will follow the same schedule.

If you can't stay for the discussion but could meet later (or earlier for Dr. Conner), let me know.

cc: Faculty



Program in Research on Women's Health

Interdisciplinary Seminar Series
Current Directions in Research on Women's Health

Cosponsored by the Breast Cancer Research Training Grant

Robert Clarke, Ph.D.

Associate Professor
Lombardi Cancer Center and
Department of Physiology & Biophysics
Georgetown University

"Estrogen Regulated Genes and Acquired Endocrine Resistance in Breast Cancer"

Date:

Thursday, March 21, 1996

Time:

4:00 - 5:00 pm

Place:

R103

Inhibition of Farnesyltransferase Induces Regression of Mammary and Salivary Carcinomas in *ras* Transgenic Mice

Dr. Michael W. Conner

Merck Research Laboratories Department of Safety

Date:

Thursday, May 23, 1996

Time:

4:00 to 5:00 p.m.

Place:

Room R103



PROGRAM IN RESEARCH ON WOMEN'S HEALTH



Issue No. 5

October 1995

Program Seminars

Michael Holick, MD, PhD, Prof. of Med., Chief, Section of Endocrinology, Diabetes & Metabolism at BU Med. Center Hospital & BCH will present a seminar entitled "New Approaches for Diagnosis and Treatment of Osteoporosis" on Wed., Nov. 8, 1995 from 3:00 to 4:00 pm in Room L-110. Refreshments will be served.

Dr. David Sherr, Prof. of Environmental Health and Pathology will present on "The role of the aromatic hydrocarbon receptor in cell growth and death" on Thursday, Dec. 7, 1995 in R-103 (the McNary Learning Cntr) from 4:00 to 5:00 pm.

Please mark your calendars for future events:

Dr. Stuart Chipkin, Assoc. Prof. of Med., and Director of Clinical Endocrinology, Diabetes and Metabolism will moderate a discussion on Obesity on Tues., Jan. 16, 1996 from 3:00 to 5:00pm. Room to be announced. As part of the Commonwealth of Massachusetts
Course on Breast Cancer,
Dr. Susan Faqua,
University of Texas
Science Center, will
present on estrogen
receptor mutations and
receptor variance in
breast cancer on Thurs.,
Feb. 22, 1996 in Atrium
C/D Conf. room at 4:00
pm. A reception follows in
Function Room A.

Dr. Robert Clarke, the Lombardi Cancer Res. Center, Georgetown, will give a seminar on Thurs., March 21, 1996 & Dr. Mike Conners of Merck Labs. will present a seminar on Thurs. May 23rd. Both lectures will be held in R103 from 4:00 to 5:00 pm.

On Thurs. June 20, 1996, **Dr. Phil Stubblefield**, Prof. and Chairman of Dept. of Obstetrics and Gyn., BU Sch. of Med., Dir., Obstetrics and Gyn., BCH, Chief, Dept. of Gynecology, BUMCH will present a seminar on **contraception.**

Seminars/Lectures

Dr. Carol Rosenberg will be presenting on "Early

Genetic Change in Breast Disease" as part of the Pathology & Lab. Med. Dept. seminar series on Fri., Nov. 17 at 1:15 pm in room L301.

Working Groups

The Breast Cancer
Working Group will
meet on Thursdays at
4:00 pm in Med/Dent Sci
conf. rm (L-317). On Oct.
26th, Mika Sovak,
MD/PhD candidate will
present her work on
"Transcription Factors in
Breast Cancer". Dr. Marc
Hansen will give the next
presentation on Nov. 30th
on "Issues and analysis
of the BRCA-1 gene".

Other News

This term's Special Topics in Path. course is on **Grant Writing**. PhD students are writing applications on aspects of research in breast cancer. Interested volunteers to review 1-3 applications for Spring term study section, please send info. on expertise, phone, and fax to Dr. Adrianne Rogers (L-804, Fax 8-4085).



PROGRAM IN RESEARCH ON WOMEN'S HEALTH

NRWSEBTOTER

Issue No. 6

February 1996

Upcoming Seminars

Suzanne Fuqua, PhD, Univ. of Texas Health Science Center will speak on "The Role of Estrogen Receptor Variants in Breast Cancer Evolution" on Thurs., Feb. 22 from 4-5pm in Atrium C/D. A reception will follow in Function Rm A. This event is co-sponsored by the Mass. Course on Breast Cancer.

Robert Clarke, PhD of the Lombardi Cancer Res. Center, will speak on: "Estrogen Regulated Genes and Acquired Endocrine Resistance in Breast Cancer" on Thurs., March 21, in R103 from 4-5pm.

Lawrence Grossman, PhD, Johns Hopkins School of Public Health, will present on "DNA Repair as a Biomarker for Cancer Epidemiology Studies". Cosponsored by the Biochem. Dept., this seminar will be held on Mon., May 13, from 4-5pm in Room L112.

Good News...

Three Breast Cancer
Working Group
members were awarded
1996 Commonwealth of
Mass. Breast Cancer
Research Awards. The
award winners were:
Ken Santora, Ph.D.
candidate (Pathology
Dept.): "Generation of
Polyclonal Antibody
Libraries for Breast
Cancer Therapy".

David Seldin, MD, PhD (Dept. of Hematology/Oncology): "Casein Kinase II in Multistep Mammary Tumorigenesis".

Mika Sovak, MD, PhD candidate (Dept. of Pathology): "Role of NF-κB/Rel in the Etiology of Breast Cancer".

Breast Cancer Working Group

The working group will hear two presentations on award-winning projects in upcoming meetings: Jackie Sharon, PhD, Dept. of Pathology & Lab. Med.

will present on:
"Polyclonal Antibody
Libraries for Cancer
Treatment", Thurs.,
Feb. 29, from 4-5pm in
room K-225.

Dave Seldin, MD, PhD, will speak on "Transgenic Mouse Models of Breast Cancer", Thurs., March 28, from 4-5pm in room K-225.

Note the room change inspired by a large turnout at the last meeting.

New Working Group

The Obesity Symposium, moderated by Dr. Stuart Chipkin attracted a grand audience with stimulating discussion. Interested P.I.'s have organized an ongoing working group which meets every other Wednesday from 10-11am in Room E210.

The next meeting is scheduled for Feb. 28th. Please contact Dr. Barbara Corkey at x8-7091 for more info.



Program in Research on Women's Health

Interdisciplinary Seminar Series
Current Directions in Research on Women's Health

Cosponsored with the Department of Biochemistry

SEMINAR

Lawrence Grossman, Ph.D.

University Distinguished Service Professor

Department of Biochemistry

The Johns Hopkins University School of Hygiene and

Public Health

DNA Repair as a Biomarker for Cancer Epidemiology Studies

Monday, May 13, 1996 4:00 p.m., Room L-112

Coffee and Tea will be served at 3:30 p.m. in the Biochemistry Conference Room (K-225)

BOSTON UNIVERSITY SCHOOL OF MEDICINE. SCHOOL OF PUBLIC HEALTH . THE UNIVERSITY HOSPITAL . BOSTON UNIVERSITY GOLDMAN SCHOOL OF GRADUATE DENTISTRY



Boston University School of Medicine

Program in Research on Women's Health Gail E. Sonenshein, Ph.D. Director Marianne Prout, M.D., M.P.H. Associate Director

80 East Concord Street Boston, Massachusetts 02118-2394 TEL: 617 638-4120 FAX: 617 638-5339

DATE:

April 24, 1996

TO:

Members of the Breast Cancer Working Group, Members of the Obesity

Group, Drs. Michael Holick and Barbara Millen

FROM:

Drs. Marianne Prout and Gail Sonenshein

RE:

The next working group meeting

The next working group meeting will be held on:

Thursday, May 2, 1996 4:00 - 5:00pm Room K-225

<u>Topic</u>: Discussion of projects which could be proposed for Program Project in Nutrition and Basic Biology Research for Cancer Prevention (RFA: CA-96-005), due July 19, 1996.

Organizer: Dr. Adrianne Rogers. Enclosed are her suggestions for focus of proposal.

Massachusetts Department of Public Health 1995-96 Breast Cancer Research Lecture Series

	Speaker	Location	Date/lime	Topic
	Dr. Mina Bissel Director, Life Sciences Lawrence Berkeley National Laboratory	Tufts Medical School Sackler Auditorium A	August 9, 1995 4:00 PM (reception follows)	"Extracellular Matrix as a Central Switch in Gene Expression Apoptosis and Cancer"
*	Dr. V. Craig Jordan, Director Breast Cancer Research Northwestern Medical School	Boston University Medical Center	September 28, 1995 4:00 PM (reception follows)	"The World Turned Upside Upside Down": "Transfection of the ER Gene Into Breast Cancer Cells"
	Dr. Paolo Toniolo Dept. of Environmental Medicine N.Y. University School of Medicine	Tufts Medical School Sackler Auditorium B	October 19, 1995 4:00 PM (reception follows)	"Endogenous Hormones in Breast Cancer
	Dr.Sara Sukumar Dept. of Oncology John's Hopkins School of Medicine	Tufts Medical School	November 29, 1995 4:00 PM (reception follows)	"Preventional Breast Cancer by Novel Gene Mediated Strategies"
	Dr. David Blask Bassett Research Institute Coopertowns, NY	Univ. of Mass. Medical Center (Grand Rounds)	January 18, 1996 12:00 noon (luncheon follows)	"The Hormone of Darkness Sheds New Light on Breast Cancer"
	Dr. Sugan Faqua (Atrec//c) Dept. of Onedlogy University of Texas Houlth Science Center	Boston Univ. Medical Center Atrium C/D	February 22, 1996 4:00 PM (reception follows) Function Room A	Estrogen Receptor Variants & Broast Cancor
义	Expert Panel (pro & con) Dr. Kathy Hetzlsouer Johns Hopkins University Dr. Marianne Prout Boston University Medical School	Harvard School of Public Health Kresge, Synder Auditorium	March 13, 1996 2:00 - 4:00 PM (reception follows) Room 110	Current Issues on Tamoxifen for Chemoprevention of Breast Cancer
·	Speaker	Location	Date/Time	Topic
\bigstar	Third Annual Massachusetts Breast Cancer Research Symposium	Gallerin Conference Center Longwood Medical Area	April 17, 1996 9:00 AM - 5:00 PM	1995 Breast Cancer Research Awardees Present Research Results
	Dr. Robert Callahan National Cancer Institute	Mass. General Hospital Cancer Center	May 8, 1996 Noon (luncheon follows)	"Genetic Alterations & Breast Cancer"
	Dr. Robert Strange Univ. of Colorado Medical Center	Mass. General Hospital	June 1996	"Apoptosis and Breast Cancer"

* students, bacusty attended

Boston University School of Medicine, 80 East Concord Street, Boston, MA 02118-2394

PATHOLOGY SEMINARS, FALL TERM 1995 FRIDAYS, 1:15-2:30 ROOM L301, REFRESHMENTS AT 1:00

September

22 Dr. John Tilley
Vincent Center for Reproductive Biology
Massachusetts General Hospital

Intracellular effectors of the ovarian cell death pathway.

29 Dr. Levon Khachigian
Department of Pathology
Brigham & Women's Hospital

Regulation of platelet derived growth factor gene expression in endothelial cells.

October

6 Dr. Daniel Wright
Department of Medicine

Basic studies of the regulation of myelopoiesis.

13 Dr. Stephen Farmer Department of Biochemistry Transcriptional control of adipogenesis and energy metabolism.

20 Kent Chen
Department of Pathology & Laboratory Medicine

Oxidative stress in atherosclerosis.

Special Seminar - Room: L110
27 Dr. Marjorie Lees
Eunice Kennedy Shriver Center
for Mental Retardation
Harvard Medical School

Myelin proteolipid protein: physiology and pathology.

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and Laboratory Medicine

Boston University School of Medicine, 80 East Concord Street, Boston, MA 02118-2394

PATHOLOGY SEMINARS, SPRING TERM 1996 FRIDAYS, 1:15-2:30 ROOM L301, REFRESHMENTS AT 1:00

FEBRUARY

9 Ken SantoraDepartment of Pathology & Laboratory Medicine

Combinatorial antibody libraries from chickens.

Dr. Shyr-Te JuDepartment of Medicine &Pathology & Laboratory Medicine

Gene regulation in activation-induced T cell death.

23 Dr. Abdulmaged Traish
Department of Biochemistry & Urology

Pathophysiology of impotence.

MARCH

1 Koren Mann Department of Pathology & Laboratory Medicine Mechanisms of polycyclic aromatic hydrocarbon-induced immunotoxicity.

8 SPRING RECESS

15 Gino DiSciullo Department of Pathology & Laboratory Medicine Studies of lymphocyte transmigration.

22 John Pak Department of Pathology & Laboratory Medicine Moloney murine leukemia virus activates NF-κB.

29 GRADUATE STUDENT RESEARCH DAY

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Department of PATHOLOGY and Laboratory Medicine

Boston University School of Medicine, 80 East Concord Street, Boston, MA 02118-2394
PATHOLOGY SEMINARS, FALL TERM 1996
FRIDAYS, 1:15-2:30
ROOM L301, REFRESHMENTS AT 1:00

September

20 Alexander Urbano

Department of Pathology & Laboratory Medicine

Identification of novel endonucleases in druginduced apoptosis of human leukemia.

27 Jennifer Cermak

Department of Pathology & Laboratory Medicine

The availability of dietary choline during pregnancy alters the brain development of offspring.

OCTOBER

11 Dr. Joseph Loscalzo

Cardiovascular Institue and Department of Medicine

Nitric oxide and homeostasis.

18 Dr. Vasilis Zannis

Departments of Medicine and Biochemistry

Apolipoprotein E and its role in the pathogenesis of cardiovascular disease and Alzheimer's disease.

20-22 Whitehead Institute Symposium at MIT

Cancer (Graduate Students encouraged to attend)

25 Ken Matsui

Department of Pathology & Laboratory Medicine

Proteasome regulation of Fas ligand.

NOVEMBER

1 Dr. David Seldin

Department of Medicine & Microbiology

Oncogenes and anti-oncogenes to model lymphomagenesis in transgenic mice.

8 Dr. James Hamilton

Department of Biophysics

Detection of crystalline cholesterol and other lipids in atherosclerotic plaques by new NMR methods.

15 Nick Husni

Department of Pathology & Laboratory Medicine

Cytokines and the diabetic fibroblast: Implications for wound healing in patients with diabetes mellitus.

Neuropeptide-related drug development against

22 Dr. Ilana Gozes

Department of Clinical Biochemistry Sackler School of Medicine

neurodegeneration.

29 HOLIDAY

DECEMBER

6 Amy Williams

Department of Pathology & Laboratory Medicine

Transcriptional regulation of platelet-derived growth factor.

13 Dr. Gerhard Heinrich

Department of Biomolecular Medicine

Zebrafish neurotrophins and trks.

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Cancer Prevention and Control Grand Rounds

Boston University Medical Center Evans Seminar Room First Floor, Evans Building* Noon—1 p.m.

*note change in location from previous year's seminars

Friday, September 29, 1995

"Can Tamoxifen Prevent Breast Cancer?"

Dr. Marianne Prout Associate Professor of Public Health and Surgery Boston University School of Medicine/ School of Public Health

Friday, October 27, 1995

"Suing the Tobacco Industry: A Good Cancer Prevention Strategy?"

Professor Richard Daynard, JD Northeastern Law School; Director, Tobacco Product Liability Project Friday, November 17, 1995

"Cancer Prevention at the Worksite: The Working Well Trial"

Dr. Glorian Sorenson Director of Community-based Research Dana Farber Cancer Institute; Associate Professor of Public Health Harvard University

Friday, December 15, 1995

"Controlling Chemotherapy-induced Nausea and Vomiting"

Dr. Paul Hesketh Chief of Medical Oncology St. Elizabeth's Medical Center; Professor of Medicine Tufts University School of Medicine

Please Post
Lunch will be provided
1.2 hours Nursing CEUs
Physician CMEs

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Cancer Prevention & Control Grand Rounds

Boston University Medical Center

Evans Seminar Room / Noon - 1 p.m.

Please Post

Friday, February 2, 1996

"The Control of Chemotherapy - Induced Nausea and Vomiting"

Dr. Paul Hesketh, Chief of Medical Oncology, St. Elizabeth's Hospital Professor of Medicine, Tufts University School of Medicine

Friday, March 1, 1996

"Can Aspirin Prevent Colorectal Cancer?"

Dr. Ed Giovanucci, Assistant Professor, Harvard Medical School

Friday, April 12, 1996

"Genetic Testing for Cancer: The Future Impact on Public Health"

Philip Reilly, MD, JD, Executive Director, Shriver Institute

Friday, May 17, 1996

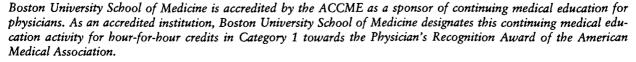
"Cigarette Advertising in Kids: Can We Compete with Joe Camel?"

Dr. Mike Siegel, Research Associate, Boston University School of Public Health

❖ Lunch will be provided ❖

1.2 hours Nursing CEUs / Physician CMEs









incer Prevention & Control Grand Rounds

Boston Medical Center

Evans Seminar Room / Evans Building / Noon - 1 p.m.

Please Post

Friday, September 27, 1996

"Special Challenges in Breast Cancer Screening and Treatment in the Inner-city"

Rita Blanchard, MD, Associate Professor of Medicine, Boston University Medical Center

Discussants:

Robert Beazley, MD, Professor of Surgery, Chief, Section of Surgical Oncology, Boston University Medical Center

Marianne Prout, MD, MPH, Associate Professor of Public Health and Surgery, Boston University Schools of Public Health and Medicine

Friday, October 18, 1996

"Vitamins and Cancer Prevention: What Should Clinicians Know?"

Donald Miller, ScD, Assistant Professor, Public Health and Dermatology, Boston University Medical Center

Discussant:

David Battinelli, MD, Director of Housestaff Education, Boston University Medical Center

Friday, November 22, 1996

"Current Approaches to Colorectal Screening"

Robert H. Fletcher, MD, MSc, Professor, Harvard Medical School

❖ Lunch will be provided ❖

1.2 hours Nursing CFUs / Physician CMEs



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CANCER

Kresge Auditorium Massachusetts Institute of Technology Cambridge, Massachusetts 1996

Sunday, October 20

KEYNOTE SPEAKERS 8:00 pm-10:00 pm

Robert A. Weinberg Richard Peto

Monday, October 21

ORIGINS

9:00 am-12:30 pm

Chairperson: Albert de la Chapelle Tom Petes Yosef Shiloh Alexander Kamb Richard Klausner

MECHANISMS

2:00 pm-5:30 pm

Chairperson: Guillermina Lozano David Beach Carol W. Greider Charles J. Sherr Eileen P. White

Tuesday, October 22

PROGRESSION

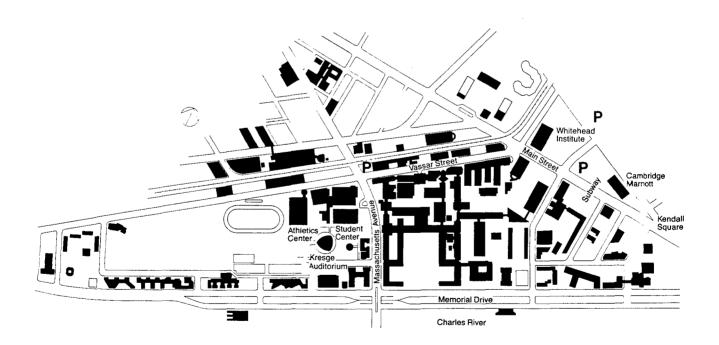
9:00 am-12:30 pm

Chairperson: Jerry Adams Douglas Hanahan Anton Berns Allan Balmain Bert Vogelstein

> **THERAPY AND** DIAGNOSTICS

2:00 pm-5:30 pm

Chairperson: Judah Folkman Allen I. Oliff Frank McCormick David Sidransky Thomas Waldmann





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Basic Rules and Policies

Research Involving Human Subjects

Leonard Glantz, J.D.

Professor of Public Health and Socio-Medical Sciences and Community Medicine, Co-Author of *Informed Consent to Human Experimentation: The Subject's Dilemma* and of *Children as Research Subjects: Science, Ethics & Law.*

The Boston University Policy on Scientific Misconduct Michael Rosen, J.D.

Associate General Counsel, Boston University; Co-Author, Boston University's Policies and Procedures Concerning Allegations of Misconduct in Scholarship and Research.

The Scientific Use of Animals: A Changing Arena

Special Guest Speaker

Peter Theran, V.M.D.

Vice-President, Hospital Division and Director of Laboratory Animal Welfare, Massachusetts Society for the Prevention of Cruelty to Animals/American Humane Education Society. Member, National Research Council Committee for Long Term Care of Chimpanzees in Research. Director of BUSM Laboratory Animal Science Center, 1966-1989.

Wednesday, September 18, 1996 3-5 pm in Room L-110

Attendance is required for pre- and post-doctoral trainees.

and please mark your calendars for..

Wednesday, October 23, 3-5pm in L-110 for a Presentation on Case Studies in Research Ethics



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RESPONSIBLE CONDUCT of RESEARCH

A Case Study on Ethical Conduct in the Lab...

You, a graduate student, are writing a paper on the regulation of IL-2 in sarcoidosis.

Your colleagues are:

A pulmonary fellow who obtained transbronchial biopsies and lung lavage fluids from the patients,

A colleague who analyzed the biopsy specimens by in situ hybridization to localize the sites of IL-2 expression,

Another colleague who has collected longitudinal epidemiologic data on the patient population of interest.

Your technician, who has performed several hundred protein assays and ELISAs for IL-2.

All contributed data and material to the paper.

- Should the technician be an author?
- Should the senior scientist in whose lab you are working be an author?
- Should the Professor of Medicine at the University of Illinois, from whom you got the highly specific IL-2 antibody be an author?
- Your epidemiologist colleague was off in Canada fly-fishing at the time you finished the manuscript and sent it off to the Journal of Biological Chemistry. She returns from her vacation, reads the accepted manuscript, and is furious because she disagrees with the conclusions. What do you do?
- What constitutes authorship?

Discussants:

Ronald Corley, Ph.D., Professor and Chairman, Microbiology Elizabeth Simons, Ph. D., Professor of Biochemistry Geoffrey Chupp, M.D., Pulmonary Fellow Caroline Fisher, M.D., Ph.D. Candidate, '99 (Pharmacology)

Moderator:

Leonard Glantz, J.D., Professor of Public Health

Wednesday, October 23, 1996 Room L-110, 3:00-5:00 p.m.

Attendance is required for pre- and post-doctoral trainees.